

Supplementary Information

Supplemental Figure Legends

Figure S1, related to Figure 1. Mouse passaged *B. thailandensis* are detected normally in vitro.

(A) IFN- γ (B-D) or LPS primed BMMs were infected with either u.p. or m.p. *Bth* for 4h. Cytotoxicity was determined by LDH release assay (A) and IL-1 β secretion, IL-1 α release, and IL-6 secretion were determined by ELISA (B-D). Bars represents mean \pm SEM of two (C-D) or three (A-B) independent experiments.

Figure S2, related to Figure 2. BALB/c mice survive *B. thailandensis* infection.

C57BL/6 or BALB/c mice were infected with the indicated doses of *B. thailandensis* and survival was monitored. Data represent one experiment with 7 mice per group.

Figure S3, related to Figure 5 and Figure 6. IFN- γ bypasses IL-18 while poly(I:C) bypasses IFN- γ .

Mice were infected with 2×10^7 cfu *B. thailandensis*. 6h later they were injected with PBS or 4 μ g IFN- γ (A) or 10mg/kg poly(I:C). 18 h later bacterial burdens were determined. Data were normalized to PBS injected controls for each organ. WT mice had no detectable bacteria (not shown). Student's *t* test (A-B).

Figure S4, related to Figure 7. Caspase-4 expression in human macrophages is largely constitutive.

Human monocyte derived macrophages were left untreated or stimulated with 1 μ g/ml LPS, 50 μ g/ml IFN- γ , or both for 24 hours. Caspase-4, pro-IL-1 β , and actin levels were determined by immunoblot.

Figure S1

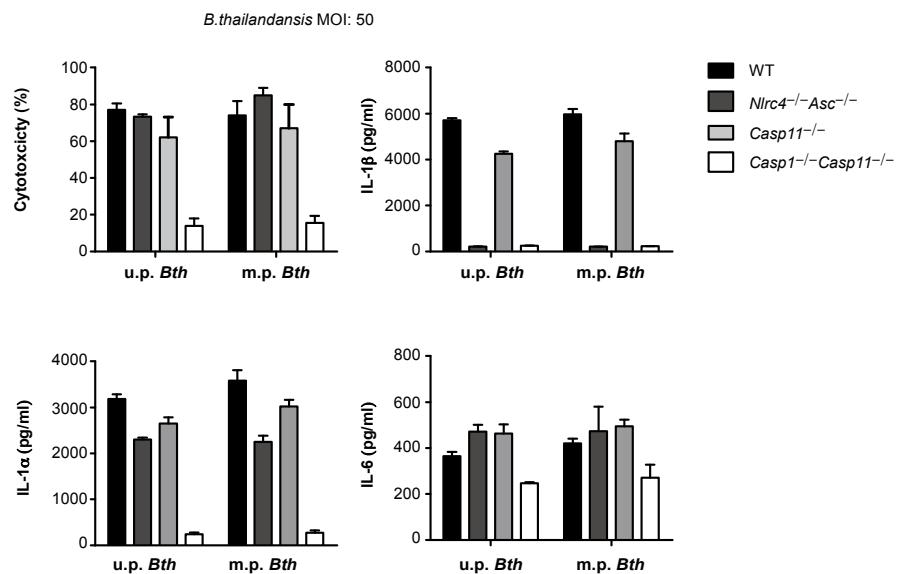


Figure S2

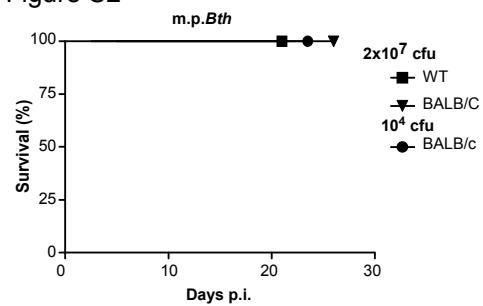


Figure S3

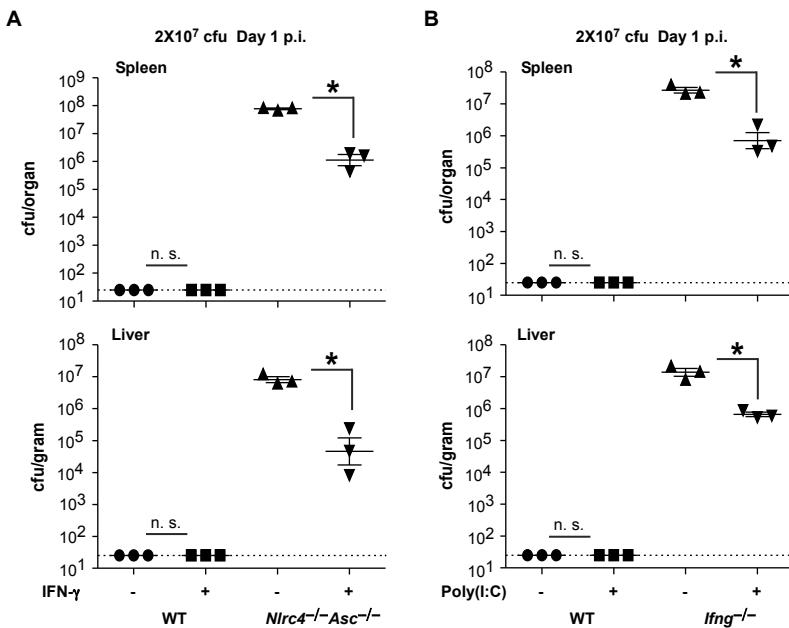
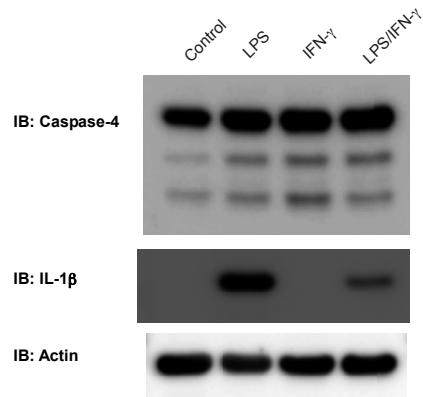


Figure S4

Casp1^{-/-} Casp11^{-/-} CASP4tg



Supplemental Tables

Table S1: related to experimental procedures. Numbers of mice used in survival experiments

Figure	Mice Genotype	n
Fig.1A	C57Bl/6 – u.p. <i>Bth</i>	5
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – u.p. <i>Bth</i>	12
	C57Bl/6– m.p. <i>Bth</i>	6
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – m.p. <i>Bth</i>	12
Fig.1B	C57Bl/6	9
	<i>Nlrc4</i> ^{-/-} <i>Asc</i> ^{-/-}	10
	<i>Casp11</i> ^{-/-}	9
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-}	9
Fig.1E	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 2x10 ⁷ cfu	6
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 10 ⁶ cfu	6
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 10 ⁵ cfu	6
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 10 ⁴ cfu	7
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 10 ³ cfu	7
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} – 10 ² cfu	7
	C57Bl/6	6
Fig.2A	<i>Nlrc4</i> ^{-/-} <i>Asc</i> ^{-/-}	12
	<i>Casp11</i> ^{-/-}	10
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-}	11
	C57Bl/6	5
Fig.2B	<i>Nlrc4</i> ^{-/-} <i>Asc</i> ^{-/-}	9
	<i>Casp11</i> ^{-/-}	12
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-}	8
	129 ^{SvEv} – 2x10 ⁷ cfu	10
Fig.2E	129 ^{SvEv} – 2.10 ⁷ cfu	10
	129 ^{SvEv} – 10 ⁴ cfu	10
	129 ^{SvEv} – 10 ² cfu	10
	C57Bl/6	4
Fig.3A	<i>Nlrc3</i> ^{-/-}	6
	<i>Nlrc5</i> ^{-/-}	6
	<i>Nlrp12</i> ^{-/-}	5
	<i>Nlrp3</i> ^{-/-} <i>Nlrp12</i> ^{-/-}	5
	<i>Nlrx1</i> ^{-/-}	9
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-}	5
	<i>Nlrp3</i> ^{-/-}	9
Fig.3D	<i>Nlrc4</i> ^{-/-}	10
	<i>Nlrp3</i> ^{-/-} <i>Nlrc4</i> ^{-/-}	9
	<i>Nlrp3</i> ^{-/-}	7
Fig.3E	<i>Nlrc4</i> ^{-/-}	9
	<i>Nlrp3</i> ^{-/-} <i>Nlrc4</i> ^{-/-}	9
	<i>Il1b</i> ^{-/-}	8
Fig.4A	<i>Il18</i> ^{-/-}	8
	<i>Il1b</i> ^{-/-} <i>Il18</i> ^{-/-}	10

	<i>Nlrc4</i> ^{-/-} <i>Asc</i> ^{-/-}	6
Fig.4B	<i>Il1b</i> ^{-/-}	9
	<i>Il18</i> ^{-/-}	8
	<i>Il1b</i> ^{-/-} <i>Il18</i> ^{-/-}	10
	<i>Nlrc4</i> ^{-/-} <i>Asc</i> ^{-/-}	5
Fig.5A	C57Bl/6	5
	<i>Ifng</i> ^{-/-}	9
	<i>Casp11</i> ^{-/-}	9
Fig.5B	C57Bl/6	5
	<i>Ifng</i> ^{-/-}	8
	<i>Casp11</i> ^{-/-}	10
Fig.5C	C57Bl/6	5
	<i>Ifng</i> ^{-/-}	9
	<i>Casp11</i> ^{-/-}	7
Fig.6A	C57Bl/6	4
	<i>Tlr4</i> ^{-/-}	6
	<i>Ifnar1</i> ^{-/-}	6
	<i>Ifngr1</i> ^{-/-} – 2x10 ⁷ cfu	7
	<i>Ifngr1</i> ^{-/-} – 10 ⁴ cfu	6
	<i>Casp11</i> ^{-/-}	5
Fig.6	C57Bl/6	5
	<i>Nos2</i> ^{-/-}	12
	<i>Casp11</i> ^{-/-}	6
Fig.7b	C57Bl/6	6
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-} <i>Casp4</i> ^{tg}	5
	<i>Casp1</i> ^{-/-} <i>Casp11</i> ^{-/-}	6